AMENDED CLAIMS

[received by the International Bureau on 10 October 2005 (10.10.2005); original claims, 16 and 17 amended; new claims 25-29 added; remaining claims unchanged (16 pages)]

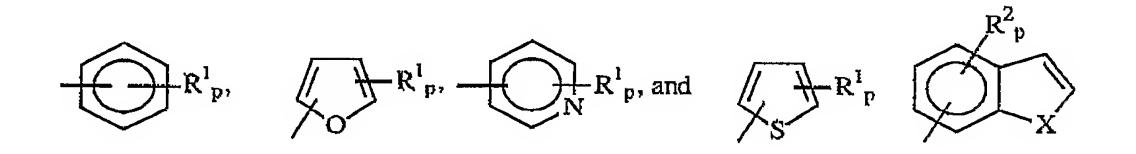
1 1. A 3,6-substituted pyran group-containing compound having

2 the structural formula:

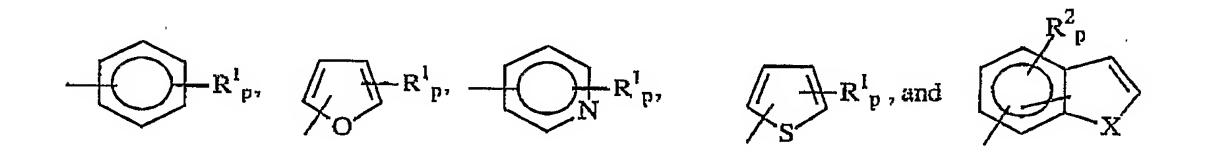
- 3 wherein
- A, A', and B are individually selected from the group of optionally substituted C_4 -
- 5 C₁₄ aryl and heteroaryl wherein heteroatoms of heteroaryl A and/or A' are selected
- from the group consisting of O, N, and S;
- Z is selected from the group consisting of a chemical bond and $-Y-(CH_{2)o}$ wherein
- 8 Y is NH or O and o is 0, 1, 2, 3, or 4;
- 9 R is H or C_{1-8} alkyl;
- W is selected from the group consisting of hydrogen and -OH; and
- n and m individually are 0, 1, 2, 3, or 4, and wherein any carbon of $-(CH_2)_n$ may
- be substituted by OR⁴ wherein R⁴ is C₁₋₈ alkyl, C₂₋₁₈ alkylene, or -COOR⁵ wherein
- 13 R^5 is C_{1-18} alkyl or C_{2-18} alkylene, and when W is H, B is an optionally substituted
- indolyl group, a 4-hydroxybenzyl group, an iodophenyl group, or a 4-aminobenzyl
- 15 group,
- or a pharmaceutically acceptable derivative or salt thereof.
- The compound of claim 1, wherein at least one of A and A' are selected from the group consisting of:

$$R^{1}_{p}$$
, R^{1}_{p} , and R^{1}_{p}

- 3 where R^1 is selected from the group consisting of C_{1-4} alkyl, C_{2-6} alkenyl, C_{2-6}
- optionally halogenated alkynyl, C₂₋₆ hydroxyalkynyl, halo, -CN, -COOR, where R
- is C₁₋₁₈ alkyl, C₅₋₁₀ cycloalkyl, C₂₋₁₈ alkenyl, -OH, -NO₂, -NH₂, -OR² where R² is
- 6 C_{1-s} alkyl, C_{5-6} cycloalkyl, or C_{2-8} alkenyl.
- 1 3. The compound of claim 1, wherein B is selected from the
- 2 group



- 3 where R^1 is selected from the group consisting of C_{1-4} alkyl, C_{2-6} alkenyl, C_{2-6}
- optionally halogenated alkynyl, C₂₋₆ hydroxyalkynyl, halo, -CN, -COOR, where R
- is C₁₋₁₈ alkyl, C₅₋₁₀ cycloalkyl, C₂₋₁₈ alkenyl, -OH, -NO₂, -NH₂, -OR² where R² is C₁₋₈
- 6 alkyl, C_{5-6} cycloalkyl, or C_{2-8} alkenyl; and
- wherein R² have the meaning of R¹ and also a 5 or 6 membered heterocycle
- 8 containing 1 or more heteroatoms selected from the group consisting of N, O, and
- 9 S, and wherein X is N, O, or S.
- 1 4. The compound of claim 2, wherein B is selected from the 2 group



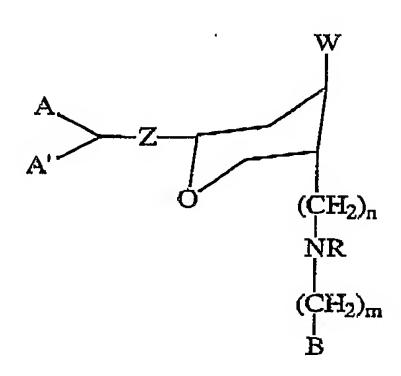
- 3 where R^1 is selected from the group consisting of $C_{1.4}$ alkyl, $C_{2.6}$ alkenyl, $C_{2.6}$
- optionally halogenated alkynyl, C₂₋₆ hydroxyalkynyl, halo, -CN, -COOR, where R

- is C_{1-18} alkyl, C_{5-10} cycloalkyl, C_{2-18} alkenyl, -OH, -NO₂, -NH₂, -OR² where R² is C_{1-8}
- 6 alkyl, C_{5-6} cycloalkyl, or C_{2-8} alkenyl; and
- wherein R² have the meaning of R¹ and also a 5 or 6 membered heterocycle
- 8 containing 1 or more heteroatoms selected from the group consisting of N, O, and
- 9 S, and wherein X is N, O, or S.
- 1 5. The compound of claim 3, wherein A and A' are both
- 2 unsubstituted phenyl.
 - 6. The compound of claim 1, having the formula

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7. The compound of claim 2, having the formula



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3

1

8. The compound of claim 3, having the formula

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1

9. The compound of claim 1, having the formula

$$A$$
 Z
 $CH_2)_n$
 NR
 $CH_2)_m$
 B

1

10. The compound of claim 2, having the formula

$$A$$
 Z
 $CH_2)_n$
 NR
 $CH_2)_m$
 B

I 11. The compound of claim 3, having the formula

$$A$$
 Z
 $CH_2)_n$
 NR
 $CH_2)_m$
 B

- 1 12. The compound of claim 1, having a formula selected from the
- 2 group consisting of:

- 1 13. The compound of claim 2, having a formula selected from the
- 2 group consisting of:

- 1 14. The compound of claim 3, having a formula selected from the
- 2 group consisting of:

$$\begin{array}{c} I \\ I \\ I \\ (CH_2)n \\ HN \\ (CH_2)m \\ R \\ \end{array}$$

$$\begin{array}{c} I \\ (CH_2)n \\ HN \\ (CH_2)m \\ R \\ \end{array}$$

$$\begin{array}{c} I \\ (CH_2)n \\ HN \\ (CH_2)m \\ R \\ \end{array}$$

$$\begin{array}{c} OH \\ (R) \\ A \\ \end{array}$$

- 1 15. The compound of claim 5, having a formula selected from the
- 2 group consisting of:

3

I

II

$$(CH_2)n$$
 $(CH_2)n$
 $(CH_2)n$
 $(CH_2)m$
 $(CH_2)m$

- 1 16. The compound of claim 1, selected from the group consisting
- 2 of:
- 3 cis-(6-benzhydryl-tetrahydropyran-3-yl)-(4-hydroxy-benzyl)-amine;
- 4 cis-(6-benzhydryl-tetrahydropyran-3-yl)-(1H-iodo-5-ylmethyl)-amine;
- 5 cis-(6-benzhydryl-tetrahydropyran-3-yl)-(4-amino-benzyl)-amine;
- 6 cis-(6-benzhydryl-tetrahydropyran-3-yl)-(3,4-dichloro-benzyl)-amine;
- 7 (2S, 4R, 5R)-2-benzhydryl-5-(4-methoxy-benzylamino)-tetrahydropyran-4-ol;
- 8 (2S, 4R, 5R)-2-benzhydryl-5-(4-fluoro-benzylamino)-tetrahydro-pyran-4-ol;
- 9 (2S, 4R, 5R)-2-benzhydryl-5-benzylamino-tetrahydro-pyran-4-ol;
- 10 (2S, 4R, 5R)-2-benzhydryl-5-(2,4-dimethoxy-benzylamino)-tetrahydropyran-4-ol;

- 11 (2S, 4R, 5R)-2-benzhydryl-5-(3,5-dimethoxy-benzylamino)-tetrahydropyran-4-ol;
- 12 (2S, 4R, 5R)-2-benzhydryi-5-(4-hydroxy-benzylamino)-tetrahydropyran-4-ol;
- 13 (2S, 4R, 5R)-2-benzhydryl-5-[(1H-indol-5-ylmethyl)-amino]-tetrahydropyran-4-ol;
- 14 (2R, 4\$, 5\$)-2-benzhydryl-5-(4-hydroxy-benzylamino)-tetrahydro-pyran-4-ol;
- 15 (2R, 4S, 5S)-2-benzhydryl-5-[(1H-indol-5-ylmethyl)-amino]-tetrahydropyran-4-ol;
- 16 cis-(3S, 6S)-(6-benzhydryl-tetrahydropyran-3-yl)-(4-hydroxy-benzyl)-amine; and
- cis-(3R, 6R)-(6-benzhydryl-tetrahydropyran-3-yl)-(4-hydroxy-benzyl)-amine.
- 1 17. The compound of claim 1, selected from the group consisting
- 2 of:
- 3 (2S, 4R, 5R)-2-benzhydryl-5-(4-methoxy-benzylamino)-tetrahydropyran-4-ol;
- 4 (2S, 4R, 5R)-2-benzhydryl-5-(4-fluoro-benzylamino)-tetrahydro-pyran-4-ol;
- 5 (2S, 4R, 5R)-2-benzhydryl-5-benzylamino-tetrahydro-pyran-4-ol;
- 6 (2S, 4R, 5R)-2-benzhydryl-5-(2,4-dimethoxy-benzylamino)-tetrahydropyran-4-ol;
- 7 (2S, 4R, 5R)-2-benzhydryl-5-(3,5-dimethoxy-benzylamino)-tetrahydropyran-4-ol;
- 8 (2S, 4R, 5R)-2-benzhydryl-5-(4-hydroxy-benzylamino)-tetrahydropyran-4-ol;
- 9 (2S, 4R, 5R)-2-benzhydryl-5-[(1H-indol-5-ylmethyl)-amino]-tetrahydropyran-4-ol;
- 10 (2R, 4S, 5S)-2-benzhydryl-5-(4-hydroxy-benzylamino)-tetrahydro-pyran-4-ol;

11 (2R, 4S, 5S)-2-benzhydryl-5-[(1H-indol-5-ylmethyl)-amino]-tetrahydropyran-4-ol; 12 cis-(3S, 6S)-(6-benzhydryl-tetrahydropyran-3-yl)-(4-hydroxy-benzyl)-amine; and 13 cis-(3R, 6R)-(6-benzhydryl-tetrahydropyran-3-yl)-(4-hydroxy-benzyl)-amine. 1 18. A method of reducing monoamine reuptake in a mammalian species, comprising administering a binding amount of a monoamine receptor binder 2 comprising a compound of claim 1. 3 A method of reducing monoamine reuptake in a mammalian 1 19. 2 species, comprising administering a binding amount of a monoamine receptor binder comprising a compound of claim 2. 3 A method of reducing monoamine reuptake in a mammalian 1 20. species, comprising administering a binding amount of a monoamine receptor binder 2 3 comprising a compound of claim 12. 1 A method for the treatment of depression, comprising 21. administering to a patient exhibiting signs of depression, a compound of claim 1 in 2 an amount effective to inhibit reuptake of serotonin at the SERT and norepinephrine 3 4 at the NET. 1 22. The method of claim 21 wherein the compound exhibits greater inhibition of serotonin and norepinephrine reuptake than of dopamine 2 3 reuptake. 1 A method for the treatment of depression, comprising 23.

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administering to a patient exhibiting signs of depression, a compound of claim 1 in

1 24. The method of claim 23 wherein said compound exhibits

- 2 higher norepinephrine reuptake inhibition than serotonin reuptake inhibition and
- 3 dopamine reuptake inhibition.
- 1 25. 1. A 3,6-substituted pyran group-containing compound
- 2 having the structural formula:

3 wherein

- A, A', and B are individually selected from the group of optionally substituted C_4
- 5 C₁₄ aryl and heteroaryl wherein heteroatoms of heteroaryl A and/or A' are selected
- from the group consisting of O, N, and S;
- 7 Z is selected from the group consisting of a chemical bond and $-Y-(CH_{2})_0$ wherein
- 8 Y is NH or O and o is 0, 1, 2, 3, or 4;
- 9 R is H or C_{1-8} alkyl;
- n and m individually are 0, 1, 2, 3, or 4, and wherein any carbon of $-(CH_2)_n$ may be
- substituted by OR⁴ wherein R⁴ is C₁₋₈ alkyl, C₂₋₁₈ alkylene, or -COOR⁵ wherein R⁵
- is C_{1-18} alkyl or C_{2-18} alkylene,
- or a pharmaceutically acceptable derivative or salt thereof.

- 1 26. The compound of claim 25, having a formula selected from
- 2 the group consisting of:

- 1 27. The compound of claim 25, having a formula selected from
- 2 the group consisting of:

3 wherein A is selected from the group consisting of:

$$R^1_p$$
, R^1_p , and R^1_p

- 4 where \mathbb{R}^1 is selected from the group consisting of \mathbb{C}_{1-4} alkyl, \mathbb{C}_{2-6} alkenyl, \mathbb{C}_{2-6}
- optionally halogenated alkynyl, C₂₋₆ hydroxyalkynyl, halo, -CN, -COOR, where R
- 6 is C_{1-18} alkyl, C_{5-10} cycloalkyl, C_{2-18} alkenyl, -OH, -NO₂, -NH₂, -OR² where R² is C_{1-8}
- 7 alkyl, C_{5-6} cycloalkyl, or C_{2-8} alkenyl.

- 1 28. The compound of claim 25,, having a formula selected from
- 2 the group consisting of:

- 3 wherein B is selected from the
- 4 group

$$\mathbb{R}^{l_p}$$
, \mathbb{R}^{l_p} , \mathbb{R}^{l_p} , and \mathbb{R}^{l_p}

- 5 where \mathbb{R}^1 is selected from the group consisting of C_{1-4} alkyl, C_{2-6} alkenyl, C_{2-6}
- optionally halogenated alkynyl, C₂₋₆ hydroxyalkynyl, halo, -CN, -COOR, where R
- is C_{1-18} alkyl, C_{5-10} cycloalkyl, C_{2-18} alkenyl, -OH, -NO₂, -NH₂, -OR² where R² is C_{1-8}
- 8 alkyl, C_{5-6} cycloalkyl, or C_{2-8} alkenyl; and

- wherein R² have the meaning of R¹ and also a 5 or 6 membered heterocycle
- containing 1 or more heteroatoms selected from the group consisting of N, O, and
- 11 S, and wherein X is N, O, or S.
- The compound of claim 27, having a formula selected form the group consisting of:

3

- 4 wherein B is selected from the
- 5 group

$$\mathbb{R}^{1}_{p}$$
, \mathbb{R}^{1}_{p} , \mathbb{R}^{1}_{p} , and \mathbb{R}^{1}_{p}

- where R^1 is selected from the group consisting of C_{1-4} alkyl, C_{2-6} alkenyl, C_{2-6}
- optionally halogenated alkynyl, C₂₋₆ hydroxyalkynyl, halo, -CN, -COOR, where R

is C_{1-18} alkyl, C_{5-10} cycloalkyl, C_{2-18} alkenyl, -OH, -NO₂, -NH₂, -OR² where R² is C_{1-8}

- 9 alkyl, C_{5-6} cycloalkyl, or C_{2-8} alkenyl; and
- 10 wherein R² have the meaning of R¹ and also a 5 or 6 membered heterocycle
- 11 containing 1 or more heteroatoms selected from the group consisting of N, O, and
- 12 S, and wherein X is N, O, or S.